-1-

SEQUENCE LISTING

	CY I	THE	CORE	PORAT	rion O	OF T	THE S	rusi	rees	OF T	THE (ORDE	ROF	THE	SISTERS	OF
<12	20>	NOVE	CL TH	ERA	PEUTI	C MC	LEC	JLES	AND	USES	THE	EREOI	ŗ			
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	0> 1>		9532 -06-													
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ctg Leu	ctc Leu	ttc Phe	tgg Trp 20	ttc Phe	ttc Phe	gat Asp	ctc Leu	gcg Ala 25	gag Glu	ccc Pro	tct Ser	ggc Gly	cgc Arg 30	gca Ala	gct Ala	96
aat Asn	gac Asp	ccc Pro 35	ttc Phe	acc Thr	atc Ile	gtc Val	cat His 40	gga Gly	aat Asn	acg Thr	ggc Gly	aag Lys 45	tgc Cys	atc Ile	aag Lys	144
cca Pro	gtg Val 50	tat Tyr	ggc Gly	tgg Trp	ata Ile	gta Val 55	gca Ala	gac Asp	gac Asp	tgt Cys	gat Asp 60	gaa Glu	act Thr	gag Glu	gac Asp	192
aag Lys 65	tta Leu	tgg Trp	aag Lys	tgg Trp	gtg Val 70	tcc Ser	cag Gln	cat His	cgg Arg	ctc Leu 75	ttt Phe	cat His	ttg Leu	cac His	tcc Ser 80	240
caa Gln	aag Lys	tgc Cys	ctt Leu	ggc Gly 85	ctc Leu	gat Asp	att Ile	acc Thr	aaa Lys 90	tcg Ser	gta Val	aat Asn	gag Glu	ctg Leu 95	aga Arg	288
atg Met	ttc Phe	agc Ser	tgt Cys 100	gac Asp	tcc Ser	agt Ser	gcc Ala	atg Met 105	ctg Leu	tgg Trp	tgg Trp	aaa Lys	tgt Cys 110	gag Glu	cac His	336

cac His	c tct s Ser	Let Let 115	TAT	gga Gly	gct Ala	gcc Ala	e cgg Arg 120	Tyr	cgg Arg	cto Lev	g gct 1 Ala	ctg Leu 125	Lуs	gat Asp	gga Gly	384
cat His	ggc Gly 130	, T11T	gca Ala	ato Ile	tca Ser	aat Asn 135	Ala	tct Ser	gat Asp	gtc Val	tgc Trp 140	Lys	aaa Lys	gga Gly	ggc Gly	432
tca Ser 145	GIU	gaa Glu	ago Ser	ctt Leu	tgt Cys 150	Asp	cag Gln	cct Pro	tat Tyr	cat His 155	Glu	atc Ile	tat Tyr	acc Thr	aga Arg 160	480
ASP	GIÀ	ASN		165	GΤĀ	Arg	Pro	Cys	Glu 170	Phe	Pro	Phe	Leu	Ile 175	Asp	528
GIY	1111	ırp	cat His 180	ніѕ	Asp	Cys	TTE	Leu 185	Asp	Glu	Asp	His	Ser 190	Gly	Pro	576
rrp	Cys	195	acc Thr	Thr	Leu	Asn	Tyr 200	Glu	Tyr	Asp	Arg	Lys 205	Trp	Gly	Ile	624
Cys	210	тÃ2	cct Pro	GIU	Asn	G1y 215	Суѕ	Glu	Asp	Asn	Trp 220	Glu	Lys	Asn	Glu	672
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БУЗ	GIU	AId	tat Tyr	245	ser	Cys	GIn	Asn	Gln 250	Gly	Ala	Asp	Leu	Leu 255	Ser	768
116	ASII	ser	gct Ala 260	Ala	GLu	Leu	Thr	Tyr 265	Leu	Lys	Glu	Lys	Glu 270	Gly	Ile	816
ALA	пуъ	275	ttc Phe	Trp	TTE	GIÀ	Leu 280	Asn	Gln	Leu	Tyr	Ser 285	Ala	Arg	Gly	864
11p	290	rrp	tca Ser	Asp	His	Lys 295	Pro	Leu	Asn	Phe	Leu 300	Asn	Trp	Asp	Pro	912
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gat Asp	gct Ala	gag Glu	tct Ser	ggt Gly 325	ctg Leu	tgg Trp	cag Gln	Ser	ttt Phe 330	tcc Ser	tgt Cys	gaa Glu	Ala	caa Gln 335	ctg Leu	1008

ecc Pro	e tai	t gte	c tgo l Cys 340	2 VI	g aaa g Lys	a cca	a tta D Lei	a aat 1 Asr 345	ı Asr	aca n Thi	a gte	g gaq l Gli	g tt: Le: 350	ı Th	a gat r Asp	1056
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aat Asn	gga Gly 370	, 1110	tgo Cys	tat Tyr	cto Leu	ctg Leu 375	ı vaı	aat . Asn	gaa Glu	agt Ser	aat Asr 380	ı Ser	tgg Trp	g gat Asp	aag Lys	1152
385	*****	, 1116	г шуз	. Cys	390	Ala	rne	ser	Ser	395	Leu	ı Ile	Ser	: Ile	cat His 400	1200
501	пец	ATO	, wah	405	GIU	vaı	vaı	Val	Thr 410	Lys	Leu	His	Asn	Glu 415		1248
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200	1110	435	пр	ser	Asp	GIÀ	440	Glu	Val	Thr	Leu	Thr 445	Tyr	Trp	gat Asp	1344
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200	273	* Y *	Val	485	ьуѕ	Arg	гуѕ	GТĀ	G1u 490	Lys	Leu	aat Asn	Asp	Ala 495	Ser	1488
		_,,0	500	Cys	110	FLO	ASP	505	стх	Trp	ЬУS	aga Arg	His 510	Gly	Glu	1536
1111	Cys	515	гуг	тте	Tyr	Glu	Asp 520	Glu	Val	Pro	Phe	gga Gly 525	Thr	Asn	Cys	1584
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aga Arg	a age	g cg g Ar	g gct g Ala 580	z val	a acc	ttt Phe	tco Ser	aac Asr 585	rrr	g aat o Asr	tti n Phe	t cti e Lei	gaq 1 Gl	ı Pr	a gct o Ala	1776
tco Sei	e cce	9 99 5 Gl; 59	A GT7	tgc Cys	gtg Val	gct Ala	atg Met 600	Ser	act Thr	: Gl	a aaq 7 Lys	g tct s Ser 605	· Val	gga Gl	a aag y Lys	1824
	610))	г пуз	, ASP	cys	615	ser	Phe	Lys	: Ala	Leu 620	ı Ser	: Ile	e Cys	c aag s Lys	1872
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705	DCI	Gly	GIII	nis	710	ren	Trp	Ile	Gly	Leu 715	Asn	aaa Lys	Arg	Ser	Pro 720	2160
		0211	Cly	725	TTD	GIN	Trp	ser	730	Arg	Thr	cca Pro	Val	Ser 735	Thr	2208
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		755	шуз	Vai	FIIG	HIS.	760	Pro	Trp	Arg	Arg	ggc Gly 765	Trp	His	Phe	2304
- 4 -	gat Asp 770	gat Asp	aga Arg	gaa Glu	FIIE	att Ile ' 775	tat Tyr	ttg Leu .	agg Arg	Pro	ttt Phe 780	gct Ala	tgt Cys	gat Asp	aca Thr	2352

aaa Lys 785		gaa u Glu	tgg Trp	g gtg Val	tgo Cys 790	GII	att Ile	cca Pro	aaa Lys	ggc Gly 795	Arc	act Thr	cca Pro	aaa Lys	aca Thr 800	2400
cca Pro	gac Asp	tgg Trp	tac Tyr	aat Asn 805	PEC	gac Asp	cgt Arg	gct Ala	gga Gly 810	' Ile	cat His	gga Gly	cct Pro	cca Pro 815	ctt Leu	2448
ata Ile	att Ile	gaa Glu	gga Gly 820	261	gaa Glu	tat Tyr	tgg Trp	ttt Phe 825	gtt Val	gct Ala	gat Asp	ctt Leu	cac His 830	cta Leu	aac Asn	2496
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	850	****	Del	rne	vaı	855	тел	ьуѕ	АТа	lle	Lys 860	Asn	aaa Lys	Ile	Ala	2592
865		501	GLY	rsp	870	GIU	тÀ2	Trp	Trp	11e 875	Arg	Ile	agc Ser	Glu	Trp 880	2640
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agc Ser	gat Asp	7117	tgt Cys 980	cac His	tcc Ser	tat Tyr	GTA (ggc Gly 985	acc Thr	ctt (Leu)	cct Pro	Ser	gtg : Val : 990	ttg a Leu S	agc Ser	2976
cag Gln	Ile	gaa Glu 995	caa (Gln)	gac i	ttt a Phe	Ile :	aca Phr 1000		ttg Leu	ctt Leu	ccg Pro	gat Asp 100	Met		a gct 1 Ala	3024

act Thr	tta Leu 1010	1-1-	att Ile	ggt Gly	ttg Leu	cgc Arg 1015	Trp	act Thr	geo Ala	tat Tyr	gaa Glu 1020	Lys	ata : Ile	aac Asn	3069
aaa Lys	tgg Trp 1025	Thr	gat Asp	aac Asn	aga Arg	gag Glu 1030	Leu	acg Thr	tac Tyr	agt Ser	aac Asn 1035	Phe		cca Pro	3114
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cct Pro	tac Tyr 1145	cag Gln	cag Gln	gca Ala	ttc Phe	ctc Leu 1150	agt Ser	gtg Val	cag Gln	gcg Ala	ctc Leu 1155	ctt Leu	cac His	aac Asn	3474
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GIU	act Thr 1190	aat Asn	GJ À aaa	caa Gln	Leu	gaa Glu 1195	gac Asp	tgt Cys	gta Val	gta Val		gac Asp			3609
GŢŽ	ttc Phe 1205	tgg Trp	aaa Lys	aca Thr	Val	gat Asp 1210	tgc Cys	aat Asn	gac Asp	Asn	caa Gln 1215	cca Pro			3654

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	123	5		· y	з су	t cca s Pro 124	0 0	r Pro	o Va.	l Le	u Asn 124	Th 5	r Pr	o Trp	
	125	0		n no	псу	t tgc s Cys 125	ту <u>г</u> 5	Ası	n Phe	∍ Il	e Ile 126	Th: O	r Ly	s Asn	
•	126	5		– 111		a cag r Gln 127(Asp)) GI	ı Val	l His	s Thr 127	Бу я 5	з Су	s Gln	
1	1280)		o ду	3 DE.	a cat r His 1285) 116	ьеи	Ser	: Ile	2 Arg 129	Asr)	Gl:	u Lys	
	1295	5		o va.	т пес	gag Glu 1300	GIN	ьeu	Leu	Туг	2 Phe 1305	Asn	tai Tyi	t atg Met	3924
N.C	1310) T.T.F	o va.	ı met	: Let	gga Gly 1315	Ile	Thr	Tyr	Arg	1320	Asn	Ser	ctt Leu	3969
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	1370	- 7 -	1116	1113	GIII	cac His 1375	ser	TIE	Leu	Ala	Cys 1380	Lys	Ile	Glu	4149
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Thr	tgg Trp 1415	tat Tyr	gaa Glu	gca Ala	tta Leu		atg (tgt Cys	tct Ser (Gln	agt Ser 1425	gga Gly	ggt Gly	cac His	4284

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	gaa Glu 1790		aca Thr	cta Leu	tgc Cys	aaa Lys 1795	aca Thr	gct Ala	atc Ile	cca Pro	tac Tyr 1800	aaa Lys	agg Arg	aaa Lys		5409
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tac Tyr	aaa Lys 1835	aaa Lys	cat His	tct Ser	Asp	tct Ser 1840	cgt Arg :	ttc Phe	acc Thr	Thr	gtt Val 1845	ttt Phe	tca Ser	acc Thr		5544

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Lys 65	Leu	Trp	Lys	Trp	Val 70	Ser	Gln	His	Arg	Leu 75	Phe	His	Leu	His	Ser 80	
Gln	Lys	Cys	Leu	Gly 85	Leu	Asp	Ile	Thr	Lys 90	Ser	Val	Asn	Glu	Leu 95	Arg	
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His	Ser	Leu 115	Tyr	Gly	Ala	Ala	Arg 120	Tyr	Arg	Leu	Ala	Leu 125	Lys	Asp	Gly	
His	Gly 130	Thr	Ala	Ile	Ser	Asn 135	Ala	Ser	Asp	Val	Trp 140	Lys	Lys	Gly	Gly	
Ser 145	Glu	Glu	Ser	Leu	Cys 150	Asp	Gln	Pro	Tyr	His 155	Glu	Ile	Tyr	Thr	Arg 160	

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Asp	Gly	Asn	Ser	Tyr 165	Gly	Arg	Pro	Cys	Glu 170	Phe	Pro	Phe	Leu	Ile 175	Asp
Gly	Thr	Trp	His 180	His	Asp	Cys	Ile	Leu 185	Asp	Glu	Asp	His	Ser 190	Gly	Pro
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Cys	Leu 210	Lys	Pro	Glu	Asn	Gly 215	Cys	Glu	Asp	Asn	Trp 220	Glu	Lys	Asn	Glu
Gln 225	Phe	Gly	Ser	Cys	Tyr 230	Gln	Phe	Asn	Thr	Gln 235	Thr	Ala	Leu	Ser	Trp 240
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Ile	Asn	Ser	Ala 260	Ala	Glu	Leu	Thr	Tyr 265	Leu	Lys	Glu	Lys	Glu 270	Gly	Ile
Ala	Lys	Ile 275	Phe	Trp	Ile	Gly	Leu 280	Asn	Gln	Leu	Tyr	Ser 285	Ala	Arg	Gly
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			Cys 340					345					350		
		355	Tyr				360					365			
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Ala 385	His	s Ala	a Lys	s Cys	390	Ala	. Phe	Ser	Ser	395		Ile	e Ser	: Ile	His 400
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Trp Glu Val Lys Asp Cys Arg Ser Phe Lys Ala Leu Ser Ile Cys Lys 615 Lys Met Ser Gly Pro Leu Gly Pro Glu Glu Ala Ser Pro Lys Pro Asp 630 Asp Pro Cys Pro Glu Gly Trp Gln Ser Phe Pro Ala Ser Leu Ser Cys 645 650 Tyr Lys Val Phe His Ala Glu Arg Ile Val Arg Lys Arg Asn Trp Glu 660 Glu Ala Glu Arg Phe Cys Gln Ala Leu Gly Ala His Leu Ser Ser Phe Ser His Val Asp Glu Ile Lys Glu Phe Leu His Phe Leu Thr Asp Gln 690 695 Phe Ser Gly Gln His Trp Leu Trp Ile Gly Leu Asn Lys Arg Ser Pro 705 Asp Leu Gln Gly Ser Trp Gln Trp Ser Asp Arg Thr Pro Val Ser Thr 730 Ile Ile Met Pro Asn Glu Phe Gln Gln Asp Tyr Asp Ile Arg Asp Cys 740 745 Ala Ala Val Lys Val Phe His Arg Pro Trp Arg Arg Gly Trp His Phe 755 760 Tyr Asp Asp Arg Glu Phe Ile Tyr Leu Arg Pro Phe Ala Cys Asp Thr 770 775 Lys Leu Glu Trp Val Cys Gln Ile Pro Lys Gly Arg Thr Pro Lys Thr 785 790 795 Pro Asp Trp Tyr Asn Pro Asp Arg Ala Gly Ile His Gly Pro Pro Leu

Ile Ile Glu Gly Ser Glu Tyr Trp Phe Val Ala Asp Leu His Leu Asn

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815

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820

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- Tyr Glu Glu Ala Val Leu Tyr Cys Ala Ser Asn His Ser Phe Leu Ala 835 840 845
- Thr Ile Thr Ser Phe Val Gly Leu Lys Ala Ile Lys Asn Lys Ile Ala 850 855 860
- Asn Ile Ser Gly Asp Gly Gln Lys Trp Trp Ile Arg Ile Ser Glu Trp 865 870 875 880
- Pro Ile Asp Asp His Phe Thr Tyr Ser Arg Tyr Pro Trp His Arg Phe 885 890 895
- Pro Val Thr Phe Gly Glu Glu Cys Leu Tyr Met Ser Ala Lys Thr Trp 900 905 910
- Leu Ile Asp Leu Gly Lys Pro Thr Asp Cys Ser Thr Lys Leu Pro Phe 915 920 925
- Ile Cys Glu Lys Tyr Asn Val Ser Ser Leu Glu Lys Tyr Ser Pro Asp 930 935 940
- Ser Ala Ala Lys Val Gln Cys Ser Glu Gln Trp Ile Pro Phe Gln Asn 945 950 955 960
- Lys Cys Phe Leu Lys Ile Lys Pro Val Ser Leu Thr Phe Ser Gln Ala 965 970 975
- Ser Asp Thr Cys His Ser Tyr Gly Gly Thr Leu Pro Ser Val Leu Ser 980 985 990
- Gln Ile Glu Gln Asp Phe Ile Thr Ser Leu Leu Pro Asp Met Glu Ala 995 1000 1005
- Thr Leu Trp Ile Gly Leu Arg Trp Thr Ala Tyr Glu Lys Ile Asn 1010 $$ 1015 $$ 1020
- Lys Trp Thr Asp Asn Arg Glu Leu Thr Tyr Ser Asn Phe His Pro 1025 1030 1035
- Leu Leu Val Ser Gly Arg Leu Arg Ile Pro Glu Asn Phe Phe Glu 1040 1040

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Leu	Tyr 1115	Lys	Ile	Ile	Pro	Lys 1120	Thr	Leu	Thr	Trp	His 1125	Ser	Ala	Lys
Arg	Glu 1130	Cys	Leu	Lys	Ser	Asn 1135	Met	Gln	Leu	Val	Ser 1140	Ile	Thr	Asp
Pro	Tyr 1145	Gln	Gln	Ala	Phe	Leu 1150	Ser	Val	Gln	Ala	Leu 1155	Leu	His	Asn
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	Glu 1580					1585					1590			
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	Gly 1655					1660	•				1665			
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Gln	Asp 1760	Asp	Asp	Glu	Asp	Leu 1765	Val	Asp	Thr	Cys	Ala 1770		Leu	His
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Ile Glu Asp Val Arg Asn Gln Cys Thr Asp His Gly Ala Asp Met Ile

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ctt Leu 145	tgc Cys	aaa Lys	aca Thr	gca Ala	atc Ile 150	cca Pro	tat Tyr	gac Asp	aag Lys	aag Lys 155	tat Tyr	tta Leu	tca Ser	gat Asp	aac Asn 160		480
cac His	att Ile	tta Leu	TTG	tcg Ser 165	act Thr	ctg Leu	gtg Val	atc Ile	gct Ala 170	agc Ser	aca Thr	gtg Val	act Thr	ctg Leu 175	gca Ala		528
gtt Val	ttg Leu	GTĀ	gcg Ala 180	gtc Val	att Ile	tgg Trp	ttc Phe	ctc Leu 185	tat Tyr	aga Arg	agg Arg	agc Ser	gca Ala 190	cgc Arg	tct Ser		576
ggc Gly	rne	acc Thr 195	tct Ser	ttc Phe	tct Ser	Pro	gca Ala 200	cca Pro	caa Gln	tca Ser	Pro	tac Tyr 205	agt Ser	gat Asp	ggc Gly		624
Cys	gct Ala 210	ctg (Leu '	gta (Val '	gtt Val	Ala	gaa Glu 215	gaa Glu	gat Asp	gaa Glu	Tyr	tct Ser 220	gtt Val	cag Gln	ctg Leu	gac Asp		672
tgag	agtt	tg g	gaac	atca	g ac	gagc	acac	tga	acac	ctt	gaca	agaa	at a	attt	cctat	•	732
gcaa	gatt	gt ca	atgta	aaaa	t tt	gcca	cgga	aaa	ctga	acc	tttt	atgg	ta t	tcct	tattc		792
															aggaa	8	352
aaaaagtagg tttcagaaaa ggaactagca cagagctaac ttacaggttt tcttaagtag									9	912							
ttttcatttg agtaaatgaa agctacagta caataaagct ggtaaaacgc aaaaaaaaaa									9	972							
aaaa	aaa															9	979

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 <212> PRT
 <213> mammalian
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 <221> misc_feature
 <222> (5)..(5)
 <223> The 'Xaa' at location 5 stands for Leu.
 <220>
 <221> misc_feature
 <222> (13)..(13)
 <223> The 'Xaa' at location 13 stands for Ala, or Val.
<220>
 <221> misc_feature
 <222> (142)..(142)
<223> The 'Xaa' at location 142 stands for Thr.
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His Glu Ala Ser Xaa Val Leu Leu Ser Leu Ala Thr Xaa Ile Phe Ala
Asp Cys Pro Ser Ser Ile Trp Val Gln Phe Gln Gly Ser Cys Tyr Thr
Phe Leu Gln Val Thr Ile Asn Val Glu Asn Ile Glu Asp Val Arg Lys
Gln Cys Thr Asp His Gly Ala Asp Leu Val Ser Ile His Asn Glu Glu
Glu Asn Ala Phe Ile Leu Asp Thr Leu Gln Lys Arg Trp Lys Gly Pro
Asp Asp Leu Leu Gly Met Phe Tyr Asp Thr Asp Asp Ala Ser Phe
                                    90
Lys Trp Phe Asp Gln Ser Asn Met Thr Phe Asp Lys Trp Ala Asp Glu
                               105
                                                   110
Asp Gly Glu Asp Leu Val Asp Thr Cys Gly Phe Leu Tyr Ala Lys Thr
                            120
Gly Glu Trp Arg Lys Gly Asn Cys Glu Met Ser Ser Val Xaa Gly Thr
    130
                       135
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Leu Cys Lys Thr Ala Ile Pro Tyr Asp Lys Lys Tyr Leu Ser Asp Asn 145 150 155 160

His Ile Leu Ile Ser Thr Leu Val Ile Ala Ser Thr Val Thr Leu Ala 165 170 175

Val Leu Gly Ala Val Ile Trp Phe Leu Tyr Arg Arg Ser Ala Arg Ser 180 185 190

Gly Phe Thr Ser Phe Ser Pro Ala Pro Gln Ser Pro Tyr Ser Asp Gly 195 200 205

Cys Ala Leu Val Val Ala Glu Glu Asp Glu Tyr Ser Val Gln Leu Asp 210 215 220

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<212> DNA

<213> mammalian

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cgtttt	tttt	tttttttt					979
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<222> <223>	, ,	(43) s any nucle:	ic acid				
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atattt	atca	gataaccgca	ttttaatatc	agctttggtg	attgctagca	cagtaattct	180
gacagt	tctg	ggagcagttg	tttggttctt	gtacaaaaga	agtttggatt	ctggtttcac	240
cacagt	tttt	tcagctgcac	accaatcacc	ttataatgat	gactgtgttt	tagtagttgc	300
agagga	aaac	gaatatgata	ttcaatttaa	ctaagatttt	ggaaatatca	gactaagaca	360
aatacc	tttc	agtgattcct	ctgtaagatt	tcaatataaa	acctgataat	gaaaattagt	420
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atg							483
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<220>
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<222> (1)..(42)
<223>
<400> 18
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Ala Ala Val Ala Asp Cys Pro Ser Ser Thr Trp Ile Gln Phe
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<2 <2	10> 11> 12> 13>	19 14 PRT mamr	nalia	ın												
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Ala 1	a Ala	a Val	L Ala	Asp 5	Cys	Pro	Ser	Ser	Thr 10	Trp) Ile	Glr	Phe	:		
<2: <2:	10> 11> 12> 13>	20 5454 DNA mamm	l nalia	n												
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ctg Leu	ctc Leu	ttc Phe	tgg Trp 20	ttc Phe	ttc Phe	gat Asp	ctc Leu	gcg Ala 25	gag	ccc Pro	tct Ser	ggc	cgc Arg 30	~~~	gct Ala	96
aat Asn	gac Asp	ccc Pro 35	ttc Phe	acc Thr	atc Ile	gtc Val	cat His 40	gga Gly	aat Asn	acg Thr	Gly	aag Lys 45	tgc Cys	atc Ile	aag Lys	144
cca Pro	gtg Val 50	tat Tyr	ggc Gly	tgg Trp	ata Ile	gta Val 55	gca Ala	gac Asp	gac Asp	tgt Cys	gat Asp 60	gaa Glu	act Thr	gag Glu	gac Asp	192
aag Lys 65	tta Leu	tgg Trp	aag Lys	tgg Trp	gtg Val 70	tcc Ser	Gln	His	cgg Arg	Leu	Phe	cat His	ttg Leu	cac His	tcc Ser 80	240
caa Gln	aag Lys	tgc Cys	ctt Leu	ggc Gly 85	ctc Leu	gat Asp	att Ile	acc Thr	aaa Lys 90	tcg Ser	gta Val	aat Asn	gag Glu	ctg Leu 95	aga Arg	288
Mec	rne	agc Ser	100	Asp	ser	Ser	Ala	Met 105	Leu	Trp	Trp	Lys	Cys 110	Glu	His	336
cac His	tct Ser	ctg Leu 115	tac Tyr	gga Gly	gct Ala	gcc Ala	cgg Arg 120	tac Tyr	cgg Arg	ctg Leu	gct Ala	ctg Leu 125	aag Lys	gat Asp	gga Gly	384

cat His	ggc Glv	aca Thr	gca Ala	ato	tca Ser	aat Asn	gca	tct	gat	gto	tgg	aag	, aaa	gga	ggc Gly	432
	130					135					140)			_	
145	GIU	GIU	ser	ьеи	150	Asp	Gln	. Pro	Туг	His 155	Glu	ı Ile	Туг	Thr	aga Arg 160	480
Asp	GTÀ	ASN	ser	165	GIY	Arg	Pro	Cys	Glu 170	Phe	Pro	Phe	Leu	11e		528
Gly ggg	acc Thr	tgg Trp	cat His 180	UTS	gat Asp	tgc Cys	att Ile	ctt Leu 185	Asp	gaa Glu	gat Asp	cat His	agt Ser 190	Gly	cca Pro	576
tgg Trp	tgt Cys	gcc Ala 195	acc Thr	acc Thr	tta Leu	aat Asn	tat Tyr 200	gaa Glu	tat Tyr	gac Asp	cga Arg	aag Lys 205	tgg Trp	ggc Gly	atc Ile	624
tgc Cys	tta Leu 210	aag Lys	cct Pro	gaa Glu	aac Asn	ggt Gly 215	tgt Cys	gaa Glu	gat Asp	aat Asn	tgg Trp 220	Glu	aag Lys	aac Asn	gag Glu	672
cag Gln 225	ttt Phe	gga Gly	agt Ser	tgc Cys	tac Tyr 230	caa Gln	ttt Phe	aat Asn	act Thr	cag Gln 235	acg Thr	gct Ala	ctt Leu	tct Ser	tgg Trp 240	720
aaa Lys	gaa Glu	gct Ala	tat Tyr	gtt Val 245	tca Ser	tgt Cys	cag Gln	aat Asn	caa Gln 250	gga Gly	gct Ala	gat Asp	tta Leu	ctg Leu 255	agc Ser	768
atc Ile	aac Asn	agt Ser	gct Ala 260	gct Ala	gaa Glu	tta Leu	act Thr	tac Tyr 265	ctt Leu	aaa Lys	gaa Glu	aaa Lys	gaa Glu 270	ggc Gly	att Ile	816
gct Ala	aag Lys	att Ile 275	ttc Phe	tgg Trp	att Ile	ggt Gly	tta Leu 280	aat Asn	cag Gln	cta Leu	tac Tyr	tct Ser 285	gct Ala	aga Arg	ggc Gly	864
tgg Trp	gaa Glu 290	tgg Trp	tca Ser	gac Asp	cac His	aaa Lys 295	cca Pro	tta Leu	aac Asn	ttt Phe	ctc Leu 300	aac Asn	tgg Trp	gat Asp	cca Pro	912
gac Asp 305	agg Arg	ccc Pro	agt Ser	gca Ala	cct Pro 310	act Thr	ata Ile	ggt Gly	ggc Gly	tcc Ser 315	agc Ser	tgt Cys	gca Ala	aga Arg	atg Met 320	960
gat Asp	gct Ala	gag Glu	tct Ser	ggt Gly 325	ctg Leu	tgg Trp	cag Gln	Ser	ttt Phe 330	tcc Ser	tgt Cys	gaa Glu	gct Ala	caa Gln 335	ctg Leu	1008
ccc Pro	tat Tyr	vaı	tgc Cys 340	agg Arg	aaa Lys	cca Pro	Leu	aat Asn 345	aat Asn	aca Thr	gtg Val	Glu	tta Leu 350	aca Thr	gat Asp	1056

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gto Val	tgg Trp	aca Thr 355	TAL	tca Ser	gat Asp	acc Thr	cgc Arg 360	tgt Cys	gat Asp	gca Ala	ggc	tgg Trp 365	Leu	cca Pro	aat Asn	11	104
aat Asn	gga Gly 370	FIIE	tgc Cys	tat Tyr	ctg Leu	ctg Leu 375	gta Val	aat Asn	gaa Glu	agt Ser	aat Asn 380	Ser	tgg Trp	gat Asp	aag Lys	11	.52
gca Ala 385	HIS	gcg Ala	aaa Lys	tgc Cys	aaa Lys 390	gcc Ala	ttc Phe	agt Ser	agt Ser	gac Asp 395	cta Leu	atc Ile	agc Ser	att Ile	cat His 400	12	200
ser	теп	Ата	gat Asp	val 405	Glu	Val	Val	Val	Thr 410	Lys	Leu	His	Asn	Glu 415	Asp	12	48
116	гуз	GIU	gaa Glu 420	Val	Trp	Ile	Gly	Leu 425	Lys	Asn	Ile	Asn	Ile 430	Pro	Thr	12	96
ьец	rne	435		Ser	Asp	GIÀ	Thr 440	Glu	Val	Thr	Leu	Thr 445	Tyr	Trp	Asp	13	44
GIU	450	GIU	cca Pro	Asn	Val	Pro 455	Tyr	Asn	Lys	Thr	Pro 460	Asn	Суѕ	Val	Ser	13	92
465	ren	GIÀ	gag Glu	Leu	470	Gln	Trp	Lys	Val	Gln 475	Ser	Cys	Glu	Glu	Lys 480	14	40
ьeu	гуз	туг	gta Val	Cys 485	гуs	Arg	Lys	Gly	Glu 490	Lys	Leu	Asn	Asp	Ala 495	Ser	14	88
ser	Asp	гуз	atg Met 500	Cys	Pro	Pro	Asp	Glu 505	Gly	Trp	Lys	Arg	His 510	Gly	Glu	15	36
THE	cys	515	aag Lys	Tie	Tyr	Glu	Asp 520	Glu	Val	Pro	Phe	Gly 525	Thr	Asn	Cys	15	84
ASN	530	Thr	atc Ile	Thr	Ser	Arg 535	Phe	Glu	Gln	Glu	Tyr 540	Leu	Asn	Asp	Leu	16:	32
545	гуѕ	гÀг	tat Tyr	Asp	Lуs 550	Ser	Leu	Arg	Lys	Tyr 555	Phe	Trp	Thr	Gly	Leu 560	168	30
aga Arg	gat Asp	gta Val	gat Asp	tct Ser 565	tgt Cys	gga Gly	gag Glu	Tyr	aac Asn 570	tgg Trp	gca Ala	act Thr	Val	ggt Gly 575	gga Gly	172	28

aga Aro	a ago g Aro	g cg	g gct g Ala 580	ı va.	a acc	ttt Phe	tcc Ser	aac Asn 585	Trp	g aat o Asn	tti Phe	t ctt e Le u	gag Glu 590	Pro	gct Ala	1776
tco Sei	e cce	g ggo 5 Gly 595	A GT?	tgo Cys	gtg Val	gct Ala	atg Met 600	Ser	act Thr	gga Gly	aaq Lys	g tot s Ser 605	Val	gga Gly	aag Lys	1824
tgg Trp	gag Glu 610	ı val	g aag L Lys	gac Asp	tgc Cys	aga Arg 615	Ser	ttc Phe	aaa Lys	gca Ala	Lev 620	ı Ser	att	tgc Cys	aag Lys	1872
625	Hec	. oer	Сіу	PIC	630	стХ	Pro	GIu	Glu	Ala 635	Ser	Pro	Lys	Pro	gat Asp 640	1920
лэр	FLO	cys	Pro	645	GTĀ	Trp	GIn	Ser	Phe 650	Pro	Ala	agt Ser	Leu	Ser 655	Cys	1968
- y -	БУЗ	Val	660	птѕ	ATA	GIU	Arg	665	Val	Arg	Lys	agg Arg	Asn 670	Trp	Glu	2016
GIU	ALG	675	Arg	rne	Cys	Gin	A1a 680	Leu	Gly	Ala	His	ctt Leu 685	Ser	Ser	Phe	2064
per	690	val	ASP	GIU	TIE	ьуs 695	GLu	Phe	Leu	His	Phe 700	tta Leu	Thr	Asp	Gln	2112
705	Der	GIY	GIII	птѕ	710	Leu	Trp	Ile	Gly	Leu 715	Asn	aaa Lys	Arg	Ser	Pro 720	2160
изр	пеп	GIII	дтХ	725	Trp	GIN	Trp	Ser	730	Arg	Thr	cca Pro	Val	Ser 735	Thr	2208
116	116	Mec	740	ASII	GIU	Pue	GIn	745	Asp	Tyr	Asp	atc Ile	Arg 750	Asp	Суз	2256
*****	7114	755	пуз	Val	rne	нтг	760	Pro	Trp	Arg	Arg	ggc Gly 765	Trp	His	Phe	2304
- , -	770	nop	nrg	GIU	rne	775	Tyr	ьeu	Arg	Pro	Phe 780	gct Ala	Cys	Asp	Thr	2352
aaa Lys 785	ctt Leu	gaa Glu	tgg Trp	vaı	tgc Cys 790	caa Gln	att Ile	cca Pro	Lys	ggc Gly 795	cgt Arg	act Thr	cca Pro	Lys	aca Thr 800	2400

cca Pro	a gad O Asp	tgg Trp	tac Tyr	aat Asn 805	Pro	gac Asp	cgt Arg	gct Ala	gga Gly 810	Ile	cat His	gga Gly	cct Pro	cca Pro 815	ctt Leu	2448
ata Ile	att : Ile	gaa Glu	gga Gly 820	agt Ser	gaa Glu	tat Tyr	tgg Trp	ttt Phe 825	Val	gct Ala	gat Asp	ctt Leu	cac His 830	cta Leu	aac Asn	2496
tat Tyr	gaa Glu	gaa Glu 835	MIG	gtc Val	ctg Leu	tac Tyr	tgt Cys 840	gcc Ala	agc Ser	aat Asn	cac His	agc Ser 845	ttt Phe	ctt Leu	gcg Ala	2544
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aat Asn 865	ata Ile	tct Ser	ggt Gly	gat Asp	gga Gly 870	cag Gln	aag Lys	tgg Trp	tgg Trp	ata Ile 875	aga Arg	att Ile	agc Ser	gag Glu	tgg Trp 880	2640
cca Pro	ata Ile	gat Asp	gat Asp	cat His 885	ttt Phe	aca Thr	tac Tyr	tca Ser	cga Arg 890	tat Tyr	cca Pro	tgg Trp	cac His	cgc Arg 895	ttt Phe	2688
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aag Lys	tgt Cys	ttt Phe	пеп	aag Lys 965	atc Ile	aaa Lys	ccc Pro	Val	tct Ser 970	ctc Leu	aca Thr	ttt Phe	tct Ser	caa Gln 975	gca Ala	2928
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cag Gln	TT6	gaa Glu 995	caa Gln	gac Asp	ttt Phe	тте .	aca Thr 1000	tcc Ser	ttg Leu	ctt Leu	ccg Pro	gat Asp 100	Me	g ga t Gl	a gct u Ala	3024
	tta Leu 1010	tgg Trp	att Ile	ggt Gly	ttg Leu	cgc Arg 101	Tr	g ac	t gce r Ala	c tai a Ty:	t gad r Gli 10:	u Ly	ag af ys I			3069

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tta Lei	ttg Leu 1040	• 4	t ag 1 Se	t gg r Gl	g ago	g ctg g Leu 1049	Arc	a ata g Ile	a cc	a ga o Gl	a aat u Asn 105	₽h€	tt Ph	t gag e Glu	3159
gaą Glu	gag Glu 1055	56	t cg	c tadg Ty:	c cad	tgt Cys 1060	Ala	cta Le	a ata ı Ile	a cto	c aac u Asn 1069	Leu	caa 1 Gl	a aaa n Lys	3204
tca Ser	ccg Pro 1070		t act e Thi	t ggg	g aco y Thr	tgg Trp 1075	Asn	ttt Phe	aca Thi	a tco Sei	c tgc Cys 1080	Ser	gaa Glu	a cgc ı Arg	3249
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agg Arg	gag Glu 1130	tgt Cys	ctg Leu	aaa Lys	agt Ser	aac Asn 1135	atg Met	cag Gln	ctg Leu	gtg Val	agc Ser 1140	Ile	acg Thr	gac Asp	3429
cct Pro	tac Tyr 1145	cag Gln	cag Gln	gca Ala	ttc Phe	ctc Leu 1150	agt Ser	gtg Val	cag Gln	gcg Ala	ctc Leu 1155	ctt Leu	cac His	aac Asn	3474
tct Ser	tcc Ser 1160	tța Leu	tgg Trp	atc Ile	gga Gly	ctc Leu 1165	ttc Phe	agt Ser	caa Gln	gat Asp	gat Asp 1170	gaa Glu	ctc Leu	aac Asn	3519
	ggt Gly 1175	tgg Trp	tca Ser	gat Asp	Gly ggg	aaa Lys 1180	cgt Arg	ctt Leu	cat His	ttt Phe	agt Ser 1185	cgc Arg	tgg Trp	gct Ala	3564
	act Thr 1190	aat Asn	G1A aaa	caa Gln	nea	gaa Glu 1195	gac Asp	tgt Cys	gta Val	gta Val	tta Leu 1200	gac Asp	act Thr	gat Asp	3609
Gly	ttc Phe 1205	tgg Trp	aaa Lys	aca Thr	Val	gat Asp 1210	tgc Cys /	aat Asn	gac Asp	aat Asn	caa Gln 1215	cca Pro	ggt Gly	gct Ala	3654
	tgc Cys 1220	tac Tyr	tat Tyr	tca Ser	GIY.	aat Asn 1225	gag a Glu :	act Thr	gaa Glu	ГÀЗ	gag Glu 1230	gtc Val	aaa Lys	cca Pro	3699

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ata Ile	cca Pro 1250	Phe	cag Gln	aac Asn	tgt Cys	tgc Cys 1255	Tyr	aat Asn	ttc Phe	ata Ile	ata : Ile 1260	Thr		aat Asn		3789
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	gca Ala 1430	agc Ser	gtt Val	cac His	aac Asn	caa Gln 1435	aat Asn	ggc Gly	cag Gln	Leu	ttt Phe 1440	ctg Leu			4	329

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cat His	t gat s Asp 1460	91.	a agi y Se:	t gaa c Glu	a tca 1 Ser	a agt Ser 1465	Phe	gaa Glu	a tgo ı Trp	g to Se:	t gat r Asp 1470	Gly	agi y Sei	t aca r Thr	,	4419
ttt Phe	gac Asp 1475	T 7 7	ato Ile	c cca e Pro	tgg Trp	aaa Lys 1480	GTĀ	caa Glr	a aca n Thr	a tci Sei	t cct r Pro 1485	Gl	a aat 7 Asr	tgt Cys		4464
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	cct Pro 1610	tca Ser	tct Ser	act Thr	Trp	att Ile 1615	cag Gln	ttc Phe	caa Gln	gac Asp	agt Ser 1620	tgt Cys	tac Tyr	att Ile	48	369
	ctc Leu 1625	caa Gln	gaa Glu	gcc Ala	Ile	aaa Lys 1630	gta Val	gaa Glu	agc Ser	ata Ile	gag Glu 1635	gat Asp			49	914
ASII	cag Gln 1640	tgt Cys	act Thr	gac Asp	His	gga Gly 1645	gcg (Ala	gac Asp	atg Met	Ile	agc Ser 1650	ata Ile			49	959

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gaa q Glu (gaa Glu 1655	gaa Glu	aat Asn	gct Ala	ttt Phe	ata Ile 1660	ьeu	gat Asp	act Thr	tto Lev	g aaa Lys 1669	Lys	caa Gln	tgg Trp	5004
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	ys . 745	ary	пуз	ıyr	ьeu	1750	Asp	Asn	His	Ile	1755	Ile	Ser	Ala	5274
	760	-10	nia	361	THE	1765	тте	Leu	Thr	Val	Leu 1770	Gly	Ala	Ile	5319
	775	. 116	beu	ıyı	туз	Lys 1780	HIS	Ser	Asp	Ser	Arg 1785	Phe	Thr	Thr	5364
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- Asn Asp Pro Phe Thr Ile Val His Gly Asn Thr Gly Lys Cys Ile Lys 35 40 45
- Pro Val Tyr Gly Trp Ile Val Ala Asp Asp Cys Asp Glu Thr Glu Asp 50 55 60
- Lys Leu Trp Lys Trp Val Ser Gln His Arg Leu Phe His Leu His Ser 65 70 75 80
- Gln Lys Cys Leu Gly Leu Asp Ile Thr Lys Ser Val Asn Glu Leu Arg 85 90 95
- Met Phe Ser Cys Asp Ser Ser Ala Met Leu Trp Trp Lys Cys Glu His 100 105 110
- His Ser Leu Tyr Gly Ala Ala Arg Tyr Arg Leu Ala Leu Lys Asp Gly 115 120 125
- His Gly Thr Ala Ile Ser Asn Ala Ser Asp Val Trp Lys Lys Gly Gly 130 135 140
- Ser Glu Glu Ser Leu Cys Asp Gln Pro Tyr His Glu Ile Tyr Thr Arg 145 150 155 160
- Asp Gly Asn Ser Tyr Gly Arg Pro Cys Glu Phe Pro Phe Leu Ile Asp 165 170 175
- Gly Thr Trp His His Asp Cys Ile Leu Asp Glu Asp His Ser Gly Pro 180 185 190
- Trp Cys Ala Thr Thr Leu Asn Tyr Glu Tyr Asp Arg Lys Trp Gly Ile 195 200 205
- Cys Leu Lys Pro Glu Asn Gly Cys Glu Asp Asn Trp Glu Lys Asn Glu 210 215 220
- Gln Phe Gly Ser Cys Tyr Gln Phe Asn Thr Gln Thr Ala Leu Ser Trp 225 230 235 240

- Lys Glu Ala Tyr Val Ser Cys Gln Asn Gln Gly Ala Asp Leu Leu Ser 245 250 255
- Ile Asn Ser Ala Ala Glu Leu Thr Tyr Leu Lys Glu Lys Glu Gly Ile 260 265 270
- Ala Lys Ile Phe Trp Ile Gly Leu Asn Gln Leu Tyr Ser Ala Arg Gly 275 280 285
- Trp Glu Trp Ser Asp His Lys Pro Leu Asn Phe Leu Asn Trp Asp Pro 290 295 300
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- Asp Ala Glu Ser Gly Leu Trp Gln Ser Phe Ser Cys Glu Ala Gln Leu 325 330 335
- Pro Tyr Val Cys Arg Lys Pro Leu Asn Asn Thr Val Glu Leu Thr Asp 340 345 350
- Val Trp Thr Tyr Ser Asp Thr Arg Cys Asp Ala Gly Trp Leu Pro Asn 355 360 365
- Asn Gly Phe Cys Tyr Leu Leu Val Asn Glu Ser Asn Ser Trp Asp Lys 370 375 380
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- Ser Leu Ala Asp Val Glu Val Val Val Thr Lys Leu His Asn Glu Asp 405 410 415
- Ile Lys Glu Glu Val Trp Ile Gly Leu Lys Asn Ile Asn Ile Pro Thr 420 425 430
- Leu Phe Gln Trp Ser Asp Gly Thr Glu Val Thr Leu Thr Tyr Trp Asp 435 440 445
- Glu Asn Glu Pro Asn Val Pro Tyr Asn Lys Thr Pro Asn Cys Val Ser 450 455 460

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685

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Sei	Ser 1160	Let)	ı Trp) Ile	Gly	Leu 1165	Phe 5	e Sei	c Glr	n Asp	Asp 1170	Glu)	Leu	a Asn
Phe	Gly 1175	Trp	Ser	: Asp	Gly	Lys 1180	Arg	j Lei	ı His	Phe	Ser 1185	Arg	Trp	Ala
Glu	Thr 1190	Asn)	Gly	Gln	Leu	Glu 1195	Asp	Cys	s Val	Val	. Leu 1200	Asp	Thr	Asp
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Met	Pro 1400	Tyr	Glu	Asp	Gly	Ile 1405	Tyr	Ser	Val	Ile	Gln 1410	Lys	Lys	Val
Thr	Trp 1415	Tyr	Glu	Ala	Leu	Asn 1420	Met	Cys	Ser	Gln	Ser 1425	Gly	Gly	His
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	1520					1525					Pro 1530			-
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Asp	Glu 1580	Asn	Lys	Phe	Val	Ser 1585	Arg	Leu	Met	Arg	Glu 1590	Asn	Asn	Asn
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Lys	Gly 1670	Pro	Asp	Asp	Ile	Leu 1675	Léu	Gly	Met	Phe	Tyr 1680	Asp	Thr	Asp
Asp	Ala 1685	Ser	Phe	Lys	Trp	Phe 1690	Asp	Asn	Ser	Asn	Met 1695	Thr	Phe	Asp
	1700					1705					Val 1710			
	1715					1720					Lys 1725			
	Val 1730					1735			_	_	Thr 1740			
Tyr	Lys 1745	Arg	ьуѕ	Tyr	Leu	Ser 1750	Asp	Asn	His	Ile	Leu 1755	Ile	Ser	Ala

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